

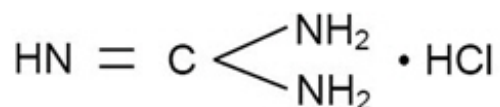
GUANIDINE HYDROCHLORIDE- guanidine hydrochloride tablet
Merck Sharp & Dohme Corp.

GUANIDINE
HYDROCHLORIDE
Tablets

DESCRIPTION

Chemically, guanidine (aminomethanamide) hydrochloride is a crystalline powder freely soluble in water and alcohol. The aqueous solution is neutral.

The structural formula is:



Each tablet contains 125 mg of guanidine hydrochloride with no color additive in the base. It also contains the following inactive ingredients: colloidal silicon dioxide, magnesium stearate, mannitol, and microcrystalline cellulose.

CLINICAL PHARMACOLOGY

Guanidine apparently acts by enhancing the release of acetylcholine following a nerve impulse. It also appears to slow the rates of depolarization and repolarization of muscle cell membranes.

INDICATIONS AND USAGE

Guanidine is indicated for the reduction of the symptoms of muscle weakness and easy fatigability associated with the myasthenic syndrome of Eaton-Lambert. It is not indicated for treating myasthenia gravis. The Eaton-Lambert syndrome is ordinarily differentiated from myasthenia gravis by the usual association of the syndrome with small cell carcinoma of the lung, but myography may be necessary to make the diagnosis.

CONTRAINDICATIONS

Guanidine is contraindicated in individuals with a history of intolerance or allergy to this drug.

WARNINGS

Fatal bone-marrow suppression, apparently dose related, can occur with guanidine.

Safe use of guanidine hydrochloride in pregnancy has not been established. Therefore, the benefits of therapy must be weighed against the potential hazards. Because guanidine is excreted in milk, patients on this drug should discontinue breastfeeding.

Since there is inadequate experience in children who have received this drug, safety and efficacy in children have not been established.

PRECAUTIONS

Baseline blood studies should be followed by frequent red and white blood cell and differential counts.

The drug should be discontinued upon appearance of bone-marrow suppression. Concurrent therapy with other drugs that may cause bone-marrow suppression should be avoided.

Renal function may be affected in some patients receiving guanidine. Patients should therefore have regular urine examinations and serum creatinine determinations while taking this drug.

Physicians should be given adequate precautions pertaining to the gastrointestinal side effects and the possibility of induced behavior disorders.

Treatment should not be continued longer than necessary.

ADVERSE REACTIONS

Anemia, leukopenia, and thrombocytopenia resulting from bone-marrow suppression attributable to guanidine have been reported. Other adverse reactions that have been observed are:

General: sore throat, rash, fever.

Neurologic: paresthesia of lips, face, hands, feet; cold sensations in hands and feet; nervousness, lightheadedness, jitteriness, increased irritability; tremor, trembling sensation; ataxia; emotional lability; psychotic state; confusion; mood changes, and hallucinations.

Gastrointestinal: dry mouth; gastric irritation; anorexia; nausea; diarrhea; abdominal cramping. Gastrointestinal side effects may preclude the use of guanidine as a desired form of therapy.

Dermatologic: rash, flushing or pink complexion; folliculitis; petechiae, purpura, ecchymoses; sweating; skin eruptions; dryness and scaling of the skin.

Renal: elevation of blood creatinine, uremia; chronic interstitial nephritis, acute interstitial nephritis, and renal tubular necrosis.

Hepatic: abnormal liver function tests.

Cardiac: palpitation, tachycardia, atrial fibrillation, hypotension.

DOSAGE AND ADMINISTRATION

Initial dosage is usually between 10 and 15 mg/kg (5 to 7 mg/pound) of body weight per day in 3 or 4 divided doses. This dosage may be gradually increased to a total daily dosage of 35 mg/kg (16 mg/pound) of body weight per day or up to the development of side effects. As individual tolerance is highly variable, the dosage must be carefully titrated. Once a tolerable dose has been established, it should be continued. Occasionally removal of the primary neoplastic lesion may result in improvement of symptoms, permitting the discontinuance of guanidine.

OVERDOSAGE

Mild gastrointestinal disorders, such as anorexia, increased peristalsis, or diarrhea are early warnings that tolerance is being exceeded. These symptoms may be relieved by atropine, but nevertheless note should be taken of these symptoms and dosage reductions considered. Slight numbness or tingling of the lips and fingertips shortly after taking a dose of guanidine has been reported. This per se is not an indication to discontinue treatment and/or reduce dosage.

Severe guanidine intoxication is characterized by nervous hyperirritability, fibrillary tremors and convulsive contractions of muscle, salivation, vomiting, diarrhea, hypoglycemia, and circulatory disturbances. Administration of intravenous calcium gluconate may control the neuromuscular and convulsive symptoms and provide some relief of other toxic manifestations.

Atropine is more effective than calcium in relieving the G.I. symptoms, circulatory disturbances, and changes in blood sugar.

HOW SUPPLIED

Guanidine hydrochloride tablets: 125 mg, white, round tablet; impressed with the product identification number "KEY 74" on one side. Guanidine hydrochloride tablets are available in bottles of 100 (NDC 0085-0492-01).

Store at 25°C (77°F); excursions permitted to 15-30°C (59-86°F) [see USP Controlled Room Temperature].

Manufactured for: Merck Sharp & Dohme Corp., a subsidiary of
MERCK & CO., INC., Whitehouse Station, NJ 08889, USA

Manufactured by: Schering Plough Canada Inc.
Pointe Claire, Quebec H9R 1B4, Canada

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O-012559-GU/HCL-TB-USPI.2
81-483544

PRINCIPAL DISPLAY PANEL - 125 mg Tablet Bottle Label

**Guanidine
Hydrochloride**

NDC 0085-0492-01

Each tablet contains

Guanidine Hydrochloride

125 mg

100 TABLETS

Rx only

Manufactured for: Merck Sharp & Dohme
Corp., a subsidiary of **Merck & Co., Inc.**,
Whitehouse Station, NJ 08889, USA

Manufactured by: Schering Plough Canada Inc.,
Pointe Claire, Quebec H9R 1B4, Canada

Formulated in Canada.

81-482245

EXP

LOT

Keep tightly closed.

NDC 0085-0492-01

Each tablet contains

Guanidine Hydrochloride

125 mg

100 TABLETS

Rx only

Manufactured for: Merck Sharp & Dohme Corp., a subsidiary of Merck & Co., Inc., Whitehouse Station, NJ 08889, USA

USUAL DOSE: See accompanying product information.

Store at 25°C (77°F); excursions permitted to 15-30°C (59-86°F) [see USP Controlled Room Temperature].

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Manufactured by: Schering Plough Canada Inc.,
Pointe Claire, Quebec H9R 1B4, Canada
Formulated in Canada.

GUANIDINE HYDROCHLORIDE

guanidine hydrochloride tablet

Product Information			
Product Type	HUMAN PRESCRIPTION DRUG	Item Code (Source)	NDC:0085-0492
Route of Administration	ORAL	DEA Schedule	
Active Ingredient/Active Moiety			
Ingredient Name		Basis of Strength	Strength
Guanidine hydrochloride (UNII: 3YQC9ZY4YB) (Guanidine - UNII:JU58 VJ6 Y3B)		Guanidine hydrochloride	125 mg
Inactive Ingredients			
Ingredient Name			Strength
silicon dioxide (UNII: ETJ7Z6XBU4)			
magnesium stearate (UNII: 70097M6I30)			
mannitol (UNII: 3OWL53L36A)			
cellulose, microcrystalline (UNII: OP1R32D61U)			
Product Characteristics			
Color	WHITE (white)	Score	no score

Shape	ROUND (round)	Size	10 mm	
Flavor		Imprint Code	KEY;74	
Contains				
Packaging				
#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0085-0492-01	100 in 1 BOTTLE		
Marketing Information				
Marketing Category	Application Number or Monograph Citation		Marketing Start Date	Marketing End Date
NDA	NDA001546		10/02/1939	

Labeler - Merck Sharp & Dohme Corp. (001317601)

Establishment			
Name	Address	ID/FEI	Business Operations
Schering Plough Canada Inc.		207093332	MANUFACTURE(0085-0492)

Establishment			
Name	Address	ID/FEI	Business Operations
Avantor Performance Materials, Inc.		152791026	API MANUFACTURE(0085-0492)